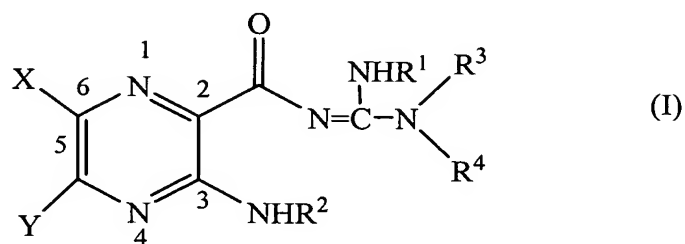


## IN THE CLAIMS

The status of each claim in the application is provided below.

Claims 1-124: Canceled.

125. (New) A compound represented by formula (I):



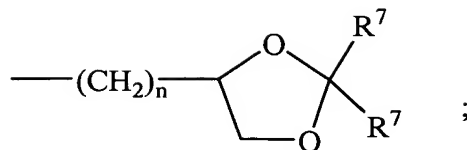
wherein

X is hydrogen, halogen, trifluoromethyl, lower alkyl, unsubstituted or substituted phenyl, lower alkyl-thio, phenyl-lower alkyl-thio, lower alkyl-sulfonyl, or phenyl-lower alkyl-sulfonyl;

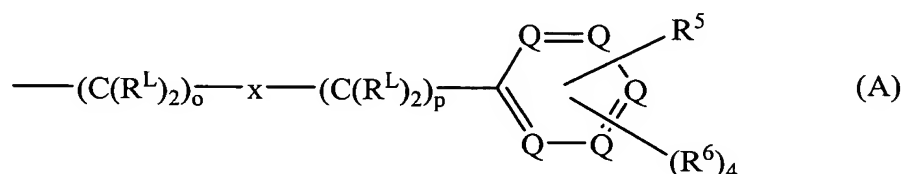
Y is hydrogen, hydroxyl, mercapto, lower alkoxy, lower alkyl-thio, halogen, lower alkyl, unsubstituted or substituted mononuclear aryl, or -N(R<sup>2</sup>)<sub>2</sub>;

R<sup>1</sup> is hydrogen or lower alkyl;

each R<sup>2</sup> is, independently, -R<sup>7</sup>, -(CH<sub>2</sub>)<sub>m</sub>-OR<sup>8</sup>, -(CH<sub>2</sub>)<sub>m</sub>-NR<sup>7</sup>R<sup>10</sup>, -(CH<sub>2</sub>)<sub>n</sub>(CHOR<sup>8</sup>)(CHOR<sup>8</sup>)<sub>n</sub>-CH<sub>2</sub>OR<sup>8</sup>, -(CH<sub>2</sub>CH<sub>2</sub>O)<sub>m</sub>-R<sup>8</sup>, -(CH<sub>2</sub>CH<sub>2</sub>O)<sub>m</sub>-CH<sub>2</sub>CH<sub>2</sub>NR<sup>7</sup>R<sup>10</sup>, -(CH<sub>2</sub>)<sub>n</sub>-C(=O)NR<sup>7</sup>R<sup>10</sup>, -(CH<sub>2</sub>)<sub>n</sub>-Z<sub>g</sub>-R<sup>7</sup>, -(CH<sub>2</sub>)<sub>m</sub>-NR<sup>10</sup>-CH<sub>2</sub>(CHOR<sup>8</sup>)(CHOR<sup>8</sup>)<sub>n</sub>-CH<sub>2</sub>OR<sup>8</sup>, -(CH<sub>2</sub>)<sub>n</sub>-CO<sub>2</sub>R<sup>7</sup>, or

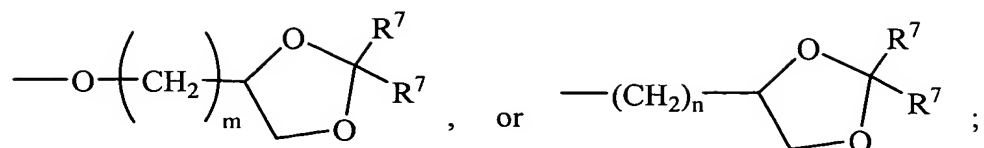


$R^3$  and  $R^4$  are each, independently, hydrogen, a group represented by formula (A), lower alkyl, hydroxy lower alkyl, phenyl, phenyl-lower alkyl, (halophenyl)-lower alkyl, lower-(alkylphenylalkyl), lower (alkoxyphenyl)-lower alkyl, naphthyl-lower alkyl, or pyridyl- lower alkyl, with the proviso that at least one of  $R^3$  and  $R^4$  is a group represented by formula (A):



wherein

each  $R^L$  is, independently,  $-R^7$ ,  $-(CH_2)_n-OR^8$ ,  $-O-(CH_2)_m-OR^8$ ,  $-(CH_2)_n-NR^7R^{10}$ ,  $-O-(CH_2)_m-NR^7R^{10}$ ,  $-(CH_2)_n(CHOR^8)(CHOR^8)_n-CH_2OR^8$ ,  $-O-(CH_2)_m(CHOR^8)(CHOR^8)_n-CH_2OR^8$ ,  $-(CH_2CH_2O)_m-R^8$ ,  $-O-(CH_2CH_2O)_m-R^8$ ,  $-(CH_2CH_2O)_m-CH_2CH_2NR^7R^{10}$ ,  $-O-(CH_2CH_2O)_m-CH_2CH_2NR^7R^{10}$ ,  $-(CH_2)_n-C(=O)NR^7R^{10}$ ,  $-O-(CH_2)_m-C(=O)NR^7R^{10}$ ,  $-(CH_2)_n-(Z)_g-R^7$ ,  $-O-(CH_2)_m-(Z)_g-R^7$ ,  $-(CH_2)_n-NR^{10}-CH_2(CHOR^8)(CHOR^8)_n-CH_2OR^8$ ,  $-O-(CH_2)_m-NR^{10}-CH_2(CHOR^8)(CHOR^8)_n-CH_2OR^8$ ,  $-(CH_2)_n-CO_2R^7$ ,  $-O-(CH_2)_m-CO_2R^7$ ,  $-OSO_3H$ ,  $-O$ -glucuronide,  $-O$ -glucose,



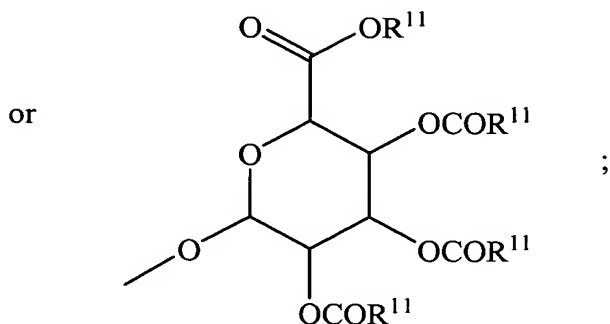
each o is, independently, an integer from 0 to 10;

each p is an integer from 0 to 10;

with the proviso that the sum of o and p in each contiguous chain is from 1 to 10;

each x is, independently, O,  $\text{NR}^{10}$ ,  $\text{C}(=\text{O})$ ,  $\text{CHOH}$ ,  $\text{C}(=\text{N}-\text{R}^{10})$ ,  $\text{CHNR}^7\text{R}^{10}$ , or represents a single bond;

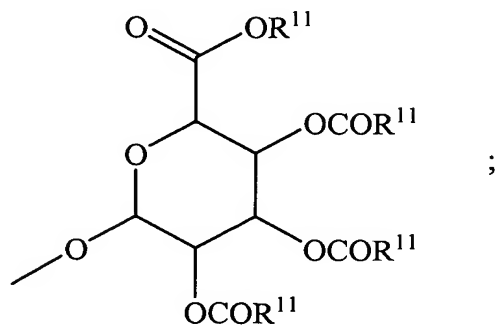
each  $\text{R}^5$  is, independently,  $-(\text{CH}_2)_m-\text{OR}^8$ ,  $-\text{O}-(\text{CH}_2)_m-\text{OR}^8$ ,  $-(\text{CH}_2)_n-\text{NR}^7\text{R}^{10}$ ,  $-\text{O}-(\text{CH}_2)_m-\text{NR}^7\text{R}^{10}$ ,  $-(\text{CH}_2)_n(\text{CHOR}^8)(\text{CHOR}^8)_n-\text{CH}_2\text{OR}^8$ ,  $-\text{O}-(\text{CH}_2)_m(\text{CHOR}^8)(\text{CHOR}^8)_n-\text{CH}_2\text{OR}^8$ ,  $-(\text{CH}_2\text{CH}_2\text{O})_m-\text{R}^8$ ,  $-\text{O}-(\text{CH}_2\text{CH}_2\text{O})_m-\text{R}^8$ ,  $-(\text{CH}_2\text{CH}_2\text{O})_m-\text{CH}_2\text{CH}_2\text{NR}^7\text{R}^{10}$ ,  $-\text{O}-(\text{CH}_2\text{CH}_2\text{O})_m-\text{CH}_2\text{CH}_2\text{NR}^7\text{R}^{10}$ ,  $-(\text{CH}_2)_n-\text{C}(=\text{O})\text{NR}^7\text{R}^{10}$ ,  $-\text{O}-(\text{CH}_2)_m-\text{C}(=\text{O})\text{NR}^7\text{R}^{10}$ ,  $-(\text{CH}_2)_n-(\text{Z})_g-\text{R}^7$ ,  $-\text{O}-(\text{CH}_2)_m-(\text{Z})_g-\text{R}^7$ ,  $-(\text{CH}_2)_n-\text{NR}^{10}-\text{CH}_2(\text{CHOR}^8)(\text{CHOR}^8)_n-\text{CH}_2\text{OR}^8$ ,  $-\text{O}-(\text{CH}_2)_m-\text{NR}^{10}-\text{CH}_2(\text{CHOR}^8)(\text{CHOR}^8)_n-\text{CH}_2\text{OR}^8$ ,  $-(\text{CH}_2)_n-\text{CO}_2\text{R}^7$ ,  $-\text{O}-(\text{CH}_2)_m-\text{CO}_2\text{R}^7$ ,  $-\text{OSO}_3\text{H}$ ,  $-\text{O-glucuronide}$ ,  $-\text{O-glucose}$ ,


$$\text{---O---}(\text{CH}_2)_m\text{---} \begin{array}{c} \text{R}^7 \\ \diagup \\ \text{O} \\ \diagdown \\ \text{R}^7 \end{array} \text{---} \quad , \quad \text{or} \quad \text{---}(\text{CH}_2)_n\text{---} \begin{array}{c} \text{R}^7 \\ \diagup \\ \text{O} \\ \diagdown \\ \text{R}^7 \end{array} \text{---} ;$$

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each  $R^7$  is, independently, hydrogen or lower alkyl;

each  $R^8$  is, independently, hydrogen, lower alkyl,  $-C(=O)-R^{11}$ , glucuronide, 2-tetrahydropyranyl, or



each  $R^9$  is, independently,  $-CO_2R^7$ ,  $-CON(R^7)_2$ ,  $-SO_2CH_3$ , or  $-C(=O)R^7$ ;

each  $R^{10}$  is, independently,  $-H$ ,  $-SO_2CH_3$ ,  $-CO_2R^7$ ,  $-C(=O)NR^7R^9$ ,  $-C(=O)R^7$ , or  $-CH_2-(CHOH)_n-CH_2OH$ ;

each  $Z$  is, independently,  $CHOH$ ,  $C(=O)$ ,  $CHNR^7R^{10}$ ,  $C=NR^{10}$ , or  $NR^{10}$ ;

each  $R^{11}$  is, independently, lower alkyl;

each  $g$  is, independently, an integer from 1 to 6;

each  $m$  is, independently, an integer from 1 to 7;

each  $n$  is, independently, an integer from 0 to 7;

each  $Q$  is, independently,  $C-R^5$ ,  $C-R^6$ , or a nitrogen atom, wherein one  $Q$  in a ring is a nitrogen atom;

or a pharmaceutically acceptable salt thereof, and

inclusive of all enantiomers, diastereomers, and racemic mixtures thereof.

126. (New) The compound of Claim 125, wherein  $Y$  is  $-NH_2$ .

127. (New) The compound of Claim 126, wherein  $R^2$  is hydrogen.
128. (New) The compound of Claim 127, wherein  $R^1$  is hydrogen.
129. (New) The compound of Claim 128, wherein X is chlorine.
130. (New) The compound of Claim 129, wherein  $R^3$  is hydrogen.
131. (New) The compound of Claim 130, wherein each  $R^L$  is hydrogen.
132. (New) The compound of Claim 131, wherein o is 4.
133. (New) The compound of Claim 132, wherein p is 0.
134. (New) The compound of Claim 133, wherein x represents a single bond.
135. (New) The compound of Claim 134, wherein each  $R^6$  is hydrogen.
136. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_m-OR^8$ .
137. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m-OR^8$ .
138. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_n-NR^7R^{10}$ .
139. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m-NR^7R^{10}$ .

140. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_n(CHOR^8)(CHOR^8)_n-CH_2OR^8$ .

141. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m(CHOR^8)(CHOR^8)_n-CH_2OR^8$ .

142. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2CH_2O)_m-R^8$ .

143. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2CH_2O)_m-R^8$ .

144. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2CH_2O)_m-CH_2CH_2NR^7R^{10}$ .

145. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2CH_2O)_m-CH_2CH_2NR^7R^{10}$ .

146. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_n-C(=O)NR^7R^{10}$ .

147. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m-C(=O)NR^7R^{10}$ .

148. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_n-(Z)_g-R^7$ .

149. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m-(Z)_g-R^7$ .

150. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_n-NR^{10}-CH_2(CHOR^8)(CHOR^8)_n-CH_2OR^8$ .

151. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m-NR^{10}-CH_2(CHOR^8)(CHOR^8)_n-CH_2OR^8$ .

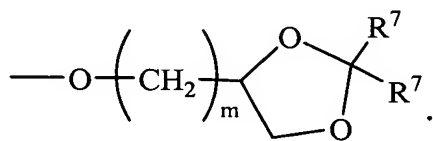
152. (New) The compound of Claim 135, wherein  $R^5$  is  $-O-(CH_2)_m-CO_2R^7$ .

153. (New) The compound of Claim 135, wherein  $R^5$  is  $-OSO_3H$ .

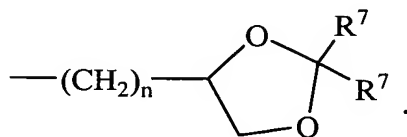
154. (New) The compound of Claim 135, wherein  $R^5$  is  $-O$ -glucuronide.

155. (New) The compound of Claim 135, wherein  $R^5$  is  $-O$ -glucose.

156. (New) The compound of Claim 135, wherein  $R^5$  is

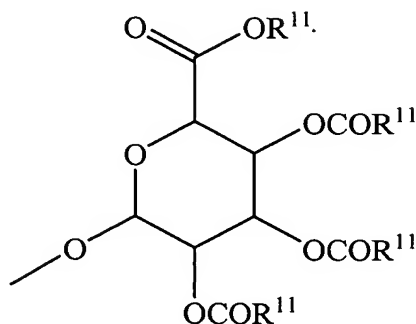


157. (New) The compound of Claim 135, wherein  $R^5$  is





158. (New) The compound of Claim 135, wherein  $R^5$  is



159. (New) The compound of Claim 135, wherein  $R^5$  is  $-(CH_2)_n-CO_2R^7$ .

160. (New) The compound of Claim 125, wherein

X is halogen;

Y is  $-N(R^7)_2$ ;

$R^1$  is hydrogen or  $C_1$ - $C_3$  alkyl;

$R^2$  is  $-R^7$ ,  $-(CH_2)_m-OR^8$ , or  $-(CH_2)_n-CO_2R^7$ ;

$R^3$  is a group represented by formula (A); and

$R^4$  is hydrogen, a group represented by formula (A), or lower alkyl.

161. (New) The compound of Claim 160, wherein

X is chloro or bromo;

Y is  $-N(R^7)_2$ ;

$R^2$  is hydrogen or  $C_1$ - $C_3$  alkyl;

at most three  $R^6$  are other than hydrogen as defined above; and

at most three  $R^L$  are other than hydrogen as defined above.

162. (New) The compound of Claim 161, wherein Y is -NH<sub>2</sub>.
163. (New) The compound of Claim 162, wherein R<sup>4</sup> is hydrogen;  
at most one R<sup>L</sup> is other than hydrogen as defined above; and  
at most two R<sup>6</sup> are other than hydrogen as defined above.
164. (New) The compound of Claim 125, wherein R<sup>5</sup> is -(CH<sub>2</sub>)<sub>m</sub>-OR<sup>8</sup>.
165. (New) The compound of Claim 125, wherein R<sup>5</sup> is -O-(CH<sub>2</sub>)<sub>m</sub>-OR<sup>8</sup>.
166. (New) The compound of Claim 125, wherein R<sup>5</sup> is -(CH<sub>2</sub>)<sub>n</sub>-NR<sup>7</sup>R<sup>10</sup>.
167. (New) The compound of Claim 125, wherein R<sup>5</sup> is -O-(CH<sub>2</sub>)<sub>m</sub>-NR<sup>7</sup>R<sup>10</sup>.
168. (New) The compound of Claim 125, wherein R<sup>5</sup> is -(CH<sub>2</sub>)<sub>n</sub>(CHOR<sup>8</sup>)(CHOR<sup>8</sup>)<sub>n</sub>-CH<sub>2</sub>OR<sup>8</sup>.
169. (New) The compound of Claim 125, wherein R<sup>5</sup> is  
-O-(CH<sub>2</sub>)<sub>m</sub>(CHOR<sup>8</sup>)(CHOR<sup>8</sup>)<sub>n</sub>-CH<sub>2</sub>OR<sup>8</sup>.
170. (New) The compound of Claim 125, wherein R<sup>5</sup> is -(CH<sub>2</sub>CH<sub>2</sub>O)<sub>m</sub>-R<sup>8</sup>.
171. (New) The compound of Claim 125, wherein R<sup>5</sup> is -O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>m</sub>-R<sup>8</sup>.

172. (New) The compound of Claim 125, wherein  $R^5$  is  $-(CH_2CH_2O)_m-CH_2CH_2NR^7R^{10}$ .

173. (New) The compound of Claim 125, wherein  $R^5$  is  $-O-(CH_2CH_2O)_m-CH_2CH_2NR^7R^{10}$ .

174. (New) The compound of Claim 125, wherein  $R^5$  is  $-(CH_2)_n-C(=O)NR^7R^{10}$ .

175. (New) The compound of Claim 125, wherein  $R^5$  is  $-O-(CH_2)_m-C(=O)NR^7R^{10}$ .

176. (New) The compound of Claim 125, wherein  $R^5$  is  $-(CH_2)_n-(Z)_g-R^7$ .

177. (New) The compound of Claim 125, wherein  $R^5$  is  $-O-(CH_2)_m-(Z)_g-R^7$ .

178. (New) The compound of Claim 125, wherein  $R^5$  is  $-(CH_2)_n-NR^{10}-CH_2(CHOR^8)(CHOR^8)_n-CH_2OR^8$ .

179. (New) The compound of Claim 125, wherein  $R^5$  is  $-O-(CH_2)_m-NR^{10}-CH_2(CHOR^8)(CHOR^8)_n-CH_2OR^8$ .

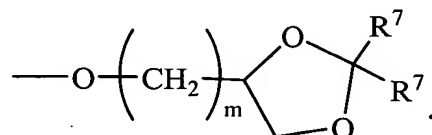
180. (New) The compound of Claim 125, wherein  $R^5$  is  $-O-(CH_2)_m-CO_2R^7$ .

181. (New) The compound of Claim 125, wherein  $R^5$  is  $-OSO_3H$ .

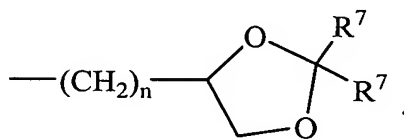
182. (New) The compound of Claim 125, wherein  $R^5$  is  $-O$ -glucuronide.

183. (New) The compound of Claim 125, wherein R<sup>5</sup> is -O-glucose.

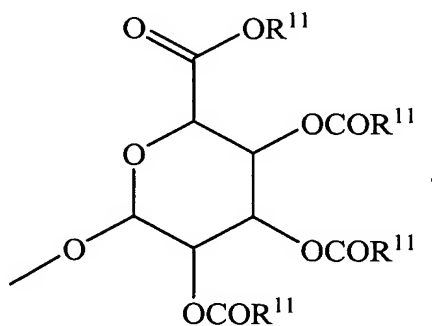
184. (New) The compound of Claim 125, wherein R<sup>5</sup> is



185. (New) The compound of Claim 125, wherein R<sup>5</sup> is



186. (New) The compound of Claim 125, wherein R<sup>5</sup> is



187. (New) The compound of Claim 125, wherein R<sup>5</sup> is  $\text{---}(\text{CH}_2)_n\text{---CO}_2\text{R}^7$ .

188. (New) The compound of Claim 125, wherein x is a single bond.

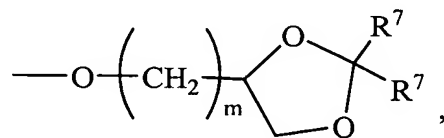
189. (New) The compound of Claim 125, which is in the form of a pharmaceutically acceptable salt.

190. (New) The compound of Claim 125, which is in the form of a hydrochloride salt.

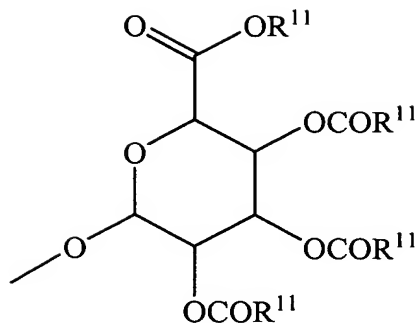
191. (New) The compound of Claim 125, which is in the form of a mesylate salt.

192. (New) The compound of Claim 125, wherein R<sup>5</sup> is selected from the group consisting of

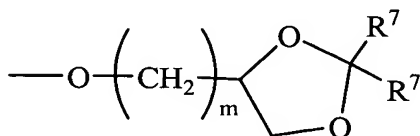
-O-(CH<sub>2</sub>)<sub>3</sub>-OH, -NH<sub>2</sub>, -O-CH<sub>2</sub>-(CHOH)<sub>2</sub>-CH<sub>2</sub>OH, -O-CH<sub>2</sub>-CHOH-CH<sub>2</sub>OH,  
-O-CH<sub>2</sub>CH<sub>2</sub>-O-tetrahydropyran-2-yl, -O-CH<sub>2</sub>CHOH-CH<sub>2</sub>-O-glucuronide,  
-O-CH<sub>2</sub>CH<sub>2</sub>OH, -O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>4</sub>-CH<sub>3</sub>, -O-CH<sub>2</sub>CH<sub>2</sub>OCH<sub>3</sub>,  
-O-CH<sub>2</sub>-(CHOC(=O)CH<sub>3</sub>)-CH<sub>2</sub>-OC(=O)CH<sub>3</sub>, -O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>2</sub>-CH<sub>3</sub>,  
-OCH<sub>2</sub>-CHOH-CHOH-CH<sub>2</sub>OH, -CH<sub>2</sub>OH, -CO<sub>2</sub>CH<sub>3</sub>,



and

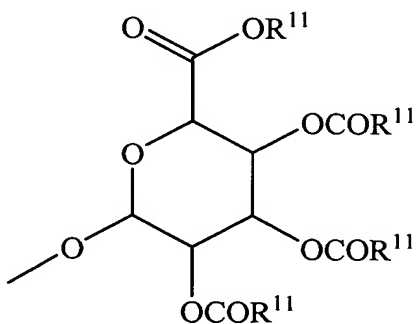


193. (New) The compound of Claim 125, wherein  $R^5$  is selected from the group consisting of para -O-(CH<sub>2</sub>)<sub>3</sub>-OH, para -NH<sub>2</sub>, para -O-CH<sub>2</sub>-(CHOH)<sub>2</sub>-CH<sub>2</sub>OH, ortho -O-CH<sub>2</sub>-CHOH-CH<sub>2</sub>OH, meta -O-CH<sub>2</sub>-CHOH-CH<sub>2</sub>OH, para -O-CH<sub>2</sub>CH<sub>2</sub>-O-tetrahydropyran-2-yl, para -O-CH<sub>2</sub>CHOH-CH<sub>2</sub>-O-glucuronide, para -O-CH<sub>2</sub>CH<sub>2</sub>OH, para -O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>4</sub>-CH<sub>3</sub>, para -O-CH<sub>2</sub>CH<sub>2</sub>OCH<sub>3</sub>, para -O-CH<sub>2</sub>-(CHOC(=O)CH<sub>3</sub>)-CH<sub>2</sub>-OC(=O)CH<sub>3</sub>, para -O-(CH<sub>2</sub>CH<sub>2</sub>O)<sub>2</sub>-CH<sub>3</sub>, -OCH<sub>2</sub>-CHOH-CHOH-CH<sub>2</sub>OH, para -CH<sub>2</sub>OH, para -CO<sub>2</sub>CH<sub>3</sub>, para -SO<sub>3</sub>H, para -O-glucuronide, para



and

para



194. (New) The compound of Claim 193, wherein

X is chloro or bromo;

Y is  $-N(R^7)_2$ ;

$R^1$  is hydrogen or  $C_1$ - $C_3$  alkyl;

$R^2$  is hydrogen or  $C_1$ - $C_3$  alkyl;

$R^3$  is a group represented by formula (A);

$R^4$  is hydrogen, a group represented by formula (A), or lower alkyl;

at most three  $R^6$  are other than hydrogen as defined above; and

at most three  $R^L$  are other than hydrogen as defined above.

195. (New) The compound of Claim 194, wherein

$R^4$  is hydrogen;

at most one  $R^L$  is other than hydrogen as defined above; and

at most two  $R^6$  are other than hydrogen as defined above.

196. (New) The compound of Claim 195, wherein

X is chloro or bromo;

Y is  $-N(R^7)_2$ ;

$R^1$  is hydrogen or  $C_1$ - $C_3$  alkyl;

$R^2$  is hydrogen or  $C_1$ - $C_3$  alkyl;

$R^3$  is a group represented by formula (A);

$R^4$  is hydrogen, a group represented by formula (A), or lower alkyl;

at most three  $R^6$  are other than hydrogen as defined above; and

at most three  $R^L$  are other than hydrogen as defined above.

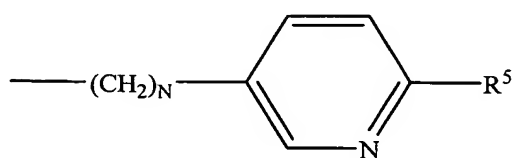
197. (New) The compound of Claim 196, wherein

$R^4$  is hydrogen;

at most one  $R^L$  is other than hydrogen as defined above; and

at most two  $R^6$  are other than hydrogen as defined above.

198. (New) The compound of Claim 125, wherein formula (A) is



wherein  $R^5$  and  $n$  are as defined in Claim 125.

199. (New) A pharmaceutical composition, comprising the compound of Claim 125 and a pharmaceutically acceptable carrier.

200. (New) A composition, comprising:

the compound of Claim 125; and

a P2Y2 inhibitor.

201. (New) A composition, comprising:

the compound of Claim 125; and

a bronchodilator.

202. (New) A method of blocking sodium channels, comprising contacting sodium channels with an effective amount of the compound of Claim 125.



### SUPPORT FOR THE AMENDMENTS

Continuing application data has been added to page 1.

The amendment at page 39 corrects an obvious error in chemical the structure of the compound over the first arrow. There should be an oxygen atom adjacent to the ClC(=O) group.

A substitute Abstract has been submitted.

Newly-added Claims 125-202 are supported by the specification at pages 4-72 and original Claims 1-124.

No new matter is believed to have been added to this application by the amendments submitted above.